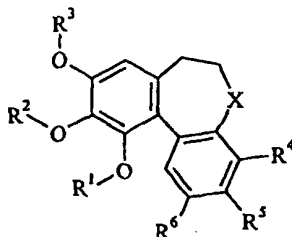


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(54) Title: COLCHINOL DERIVATIVES AS VASCULAR DAMAGING AGENTS



(I)

(57) Abstract

The invention relates to the use of compounds of formula (I): wherein X is $-\text{C}(\text{O})-$, $-\text{C}(\text{S})-$, $-\text{C}=\text{NOH}$, or $-\text{CH}(\text{R}^7)-$ wherein R^7 is hydrogen, hydroxy, C_{1-7} alkoxy, $-\text{OR}^8$ or $-\text{NR}^8\text{R}^9$ (wherein R^8 is a group $-\text{Y}^1\text{R}^{10}$ (wherein Y^1 is a direct bond, $-\text{C}(\text{O})-$, $-\text{C}(\text{S})-$, $-\text{S}-$, $-\text{C}(\text{O})\text{O}$, $-\text{C}(\text{O})\text{NR}^{11}-$, $-\text{SO}_2-$ or $-\text{SO}_2\text{NR}^{12}-$ (wherein R^{11} and R^{12} , which may be the same or different, each independently represents hydrogen, C_{1-3} alkyl or C_{1-3} alkoxy C_{2-3} alkyl) and R^{10} is as defined herein, and R^9 includes hydrogen; R^1 , R^2 and R^3 are as defined herein and are preferably methyl; R^4 , R^5 and R^6 are as defined herein with the proviso that R^5 is not hydroxy, alkoxy, substituted alkoxy, $-\text{OPO}_3\text{H}_2$, $-\text{O}-\text{C}_{1-7}$ alkanoyl or benzyloxy; and salts thereof in the manufacture of a medicament for use in the production of a vascular damaging effect in warm-blooded animals such as humans. The present invention further relates to compounds of the formula (I), pharmaceutical compositions containing them, processes for their preparation and to a method of treatment using the compounds to produce a vascular damaging effect in a warm-blooded animal such as a human. The compounds of formula (I) and the pharmaceutically acceptable salts thereof may be useful in the treatment of a number of disease states including cancer and rheumatoid arthritis.

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